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* * * * * Welcome to STN International * * * * *

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NEWS 2 NOV 21 CAS patent coverage to include exemplified prophetic
substances identified in English-, French-, German-,
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NEWS 3 NOV 26 MARPAT enhanced with FSORT command
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NEWS 5 NOV 26 Two new SET commands increase convenience of STN
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NEWS 7 DEC 12 GBFULL now offers single source for full-text
coverage of complete UK patent families
NEWS 8 DEC 17 Fifty-one pharmaceutical ingredients added to PS
NEWS 9 JAN 06 The retention policy for unread STNmail messages
will change in 2009 for STN-Columbus and STN-Tokyo
NEWS 10 JAN 07 WPIDS, WPINDEX, and WPIX enhanced Japanese Patent
Classification Data

NEWS EXPRESS JUNE 27 08 CURRENT WINDOWS VERSION IS V8.3,
AND CURRENT DISCOVER FILE IS DATED 23 JUNE 2008.

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* * * * * STN Columbus * * * * *

FILE 'HOME' ENTERED AT 09:49:04 ON 21 JAN 2009

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FILE 'REGISTRY' ENTERED AT 09:49:16 ON 21 JAN 2009

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STRUCTURE FILE UPDATES: 20 JAN 2009 HIGHEST RN 1094597-78-0
DICTIONARY FILE UPDATES: 20 JAN 2009 HIGHEST RN 1094597-78-0

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH July 5, 2008.

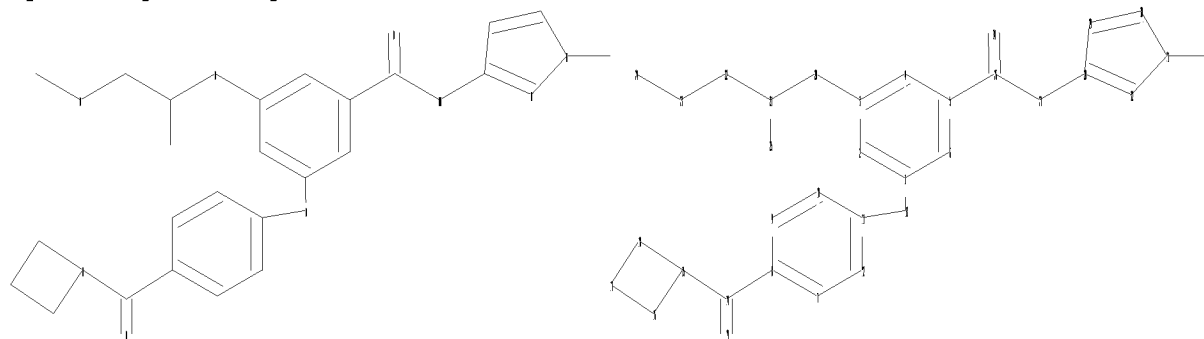
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REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

<http://www.cas.org/support/stngen/stndoc/properties.html>

=>

Uploading C:\Program Files\STNEXP\Queries\10588334 elected.str



chain nodes :

13 14 19 20 21 22 23 24 25 26 27 33 34

ring nodes :

1 2 3 4 5 6 7 8 9 10 11 12 15 16 17 18 28 29 30 31 32

chain bonds :

1-13 3-20 5-26 8-14 11-13 14-15 14-19 20-21 21-22 21-25 22-23 23-24
26-27 26-34 27-28 31-33

ring bonds :

1-2 1-6 2-3 3-4 4-5 5-6 7-8 7-12 8-9 9-10 10-11 11-12 15-16 15-18
16-17 17-18 28-29 28-32 29-30 30-31 31-32

exact/norm bonds :

1-13 3-20 11-13 14-15 14-19 15-16 15-18 16-17 17-18 20-21 22-23 23-24
26-27 26-34 27-28 28-29 28-32 29-30 30-31 31-32 31-33

exact bonds :

5-26 8-14 21-22 21-25

normalized bonds :

1-2 1-6 2-3 3-4 4-5 5-6 7-8 7-12 8-9 9-10 10-11 11-12

Match level :

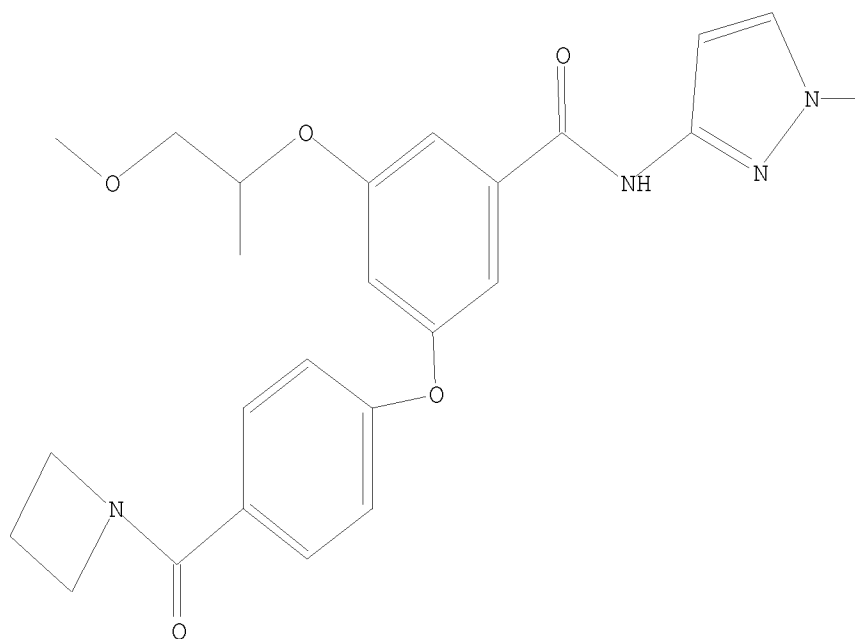
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom
11:Atom 12:Atom 13:CLASS 14:CLASS 15:Atom 16:Atom 17:Atom 18:Atom 19:CLASS
20:CLASS 21:CLASS 22:CLASS 23:CLASS 24:CLASS 25:CLASS 26:CLASS 27:CLASS
28:Atom 29:Atom 30:Atom 31:Atom 32:Atom 33:CLASS 34:CLASS

L1 STRUCTURE UPLOADED

=> d l1

L1 HAS NO ANSWERS

L1 STR



Structure attributes must be viewed using STN Express query preparation.

=> s l1 sss sam

SAMPLE SEARCH INITIATED 09:49:46 FILE 'REGISTRY'

SAMPLE SCREEN SEARCH COMPLETED - 9 TO ITERATE

100.0% PROCESSED 9 ITERATIONS

1 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**
BATCH **COMPLETE**

PROJECTED ITERATIONS: 9 TO 360

PROJECTED ANSWERS: 1 TO 80

L2 1 SEA SSS SAM L1

=> s l1 sss ful

FULL SEARCH INITIATED 09:49:51 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 121 TO ITERATE

100.0% PROCESSED 121 ITERATIONS

9 ANSWERS

SEARCH TIME: 00.00.01

L3 9 SEA SSS FUL L1

=> fil cap

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

ENTRY

SESSION

FULL ESTIMATED COST

185.88

186.10

FILE 'CAPLUS' ENTERED AT 09:49:55 ON 21 JAN 2009

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FILE COVERS 1907 - 21 Jan 2009 VOL 150 ISS 4

FILE LAST UPDATED: 20 Jan 2009 (20090120/ED)

Caplus now includes complete International Patent Classification (IPC) reclassification data for the third quarter of 2008.

Effective October 17, 2005, revised CAS Information Use Policies apply. They are available for your review at:

<http://www.cas.org/legal/infopolicy.html>

=> d 13

YOU HAVE REQUESTED DATA FROM FILE 'REGISTRY' - CONTINUE? (Y)/N:n

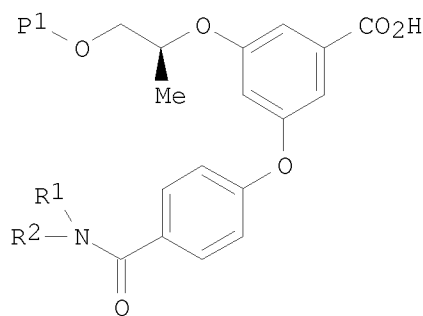
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L4 4 L3

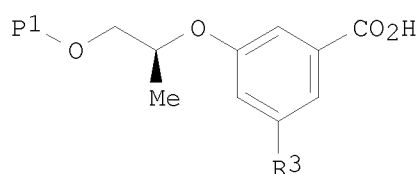
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L4 ANSWER 1 OF 4 CAPLUS COPYRIGHT 2009 ACS on STN

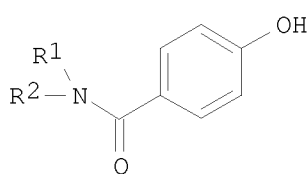
GI



I



II



III

AB The present invention relates to an improved process for preparing I [R1 and R2 independently = H, alkyl; R1 and R2 may join together with N to form a 4- to 7-membered heterocycllyl ring; P1 = H or hydroxy protecting group], which are useful as intermediates to compds. which activate glucokinase. The method involves substitution of II [R3 = halo] with III to provide I in the presence of copper catalysts containing 2,2,6,6-tetramethylheptane-3,5-dione (TMHD) ligand. An alternative preparation of I via hydrolysis of corresponding ester is provided. I could be further reacted with (un)substituted 5 to 6-membered heterocyclic amine derivs. to generate corresponding amides by amidation. Thus, e.g., substitution of 3-bromo-5-((1S)-2-tert-butoxy-1-methylethoxy)benzoic acid (preparation given) with 4-(azetidin-1-ylcarbonyl)phenol (preparation given) using

copper iodide as catalyst and TMHD as ligand gives desired intermediate 3-[4-(azetidin-1-ylcarbonyl)phenoxy]-5-[(1S)-2-tert-butoxy-1-methylethoxy]benzoic acid, which could further react with 1-methyl-3-aminopyrazole followed by hydrolysis to provide 3-[4-(azetidin-1-ylcarbonyl)phenoxy]-5-[(1S)-2-hydroxy-1-methylethoxy]-N-(1-methyl-1H-pyrazol-3-yl)benzamide.

ACCESSION NUMBER: 2007:593421 CAPLUS

DOCUMENT NUMBER: 147:30824

TITLE: Method for preparing benzoic acid derivatives via substitution of haloalkoxybenzoic acid with aminocarbonylphenol utilizing copper catalysts containing 2,2,6,6-tetramethylheptane-3,5-dione ligand
INVENTOR(S): Hopes, Phillip Anthony; Parker, Jeremy Stephen; Patel, Bharti; Welham, Matthew James

PATENT ASSIGNEE(S): Astrazeneca AB, Swed.; Astrazeneca UK Limited

SOURCE: PCT Int. Appl., 29pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.

KIND

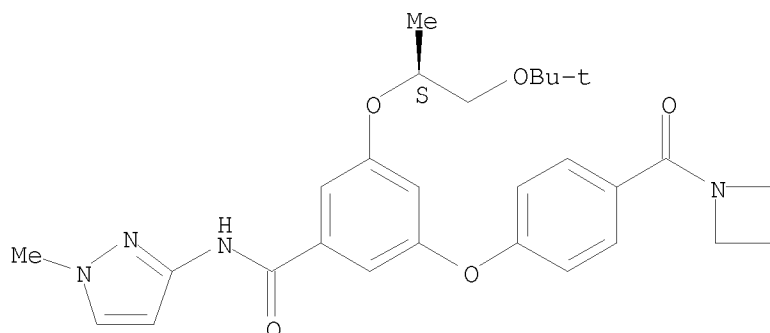
DATE

APPLICATION NO.

DATE

| | | | | |
|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----|----------|------------------|------------|
| WO 2007060448 | A2 | 20070531 | WO 2006-GB4399 | 20061127 |
| WO 2007060448 | A3 | 20080410 | | |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, GT, HN, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LA, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MY, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RS, RU, SC, SD, SE, SG, SK, SL, SM, SV, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, ZA, ZM, ZW | | | | |
| RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AP, EA, EP, OA | | | | |
| AU 2006318889 | A1 | 20070531 | AU 2006-318889 | 20061127 |
| CA 2629995 | A1 | 20070531 | CA 2006-2629995 | 20061127 |
| EP 1960354 | A2 | 20080827 | EP 2006-808668 | 20061127 |
| R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, AL, BA, HR, MK, RS | | | | |
| CN 101316815 | A | 20081203 | CN 2006-80044216 | 20080526 |
| MX 200806788 | A | 20080604 | MX 2008-6788 | 20080527 |
| IN 2008DN04513 | A | 20080815 | IN 2008-DN4513 | 20080527 |
| US 20080300412 | A1 | 20081204 | US 2008-95101 | 20080527 |
| KR 2008072738 | A | 20080806 | KR 2008-715241 | 20080623 |
| PRIORITY APPLN. INFO.: | | | US 2005-740042P | P 20051128 |
| | | | WO 2006-GB4399 | W 20061127 |
| OTHER SOURCE(S): MARPAT 147:30824 | | | | |
| IT 937842-59-6P | | | | |
| RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) | | | | |
| (preparation of aryl amide using intermediate aminocarbonylphenoxyalkoxybenzoic acid with heterocyclic amine derivs. by amidation) | | | | |
| RN 937842-59-6 CAPLUS | | | | |
| CN Benzamide, 3-[4-(1-azetidinyllcarbonyl)phenoxy]-5-[(1S)-2-(1,1-dimethylethoxy)-1-methylethoxy]-N-(1-methyl-1H-pyrazol-3-yl)- (CA INDEX NAME) | | | | |

Absolute stereochemistry.



* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT *

AB Title compds. I [R1 = fluoromethoxymethyl, difluoromethoxymethyl or trifluoromethoxymethyl; R2 = -C(O)NR4R5, -SO2NR4R5, -S(O)pR4, etc.; Het-1 = 5- or 6-membered, C-linked heteroaryl ring containing a nitrogen atom in the 2-position and optionally 1 or 2 further ring heteroatoms selected from O, N and S, which ring is optionally substituted on an available carbon atom, or on a ring nitrogen atom provided it is not thereby quaternized, with 1 or 2 substituents selected from R6; R3 = halo; R4 = H, alkyl [optionally substituted by 1 or 2 substituents selected from -OR5, -SO2R5, cycloalkyl (optionally substituted by R7)], etc.; R5 = H, alkyl; R4 and R5 together with the nitrogen atom to which they are attached may form a heterocycle; R6 = alkyl, hydroxyalkyl, alkocyalkyl, etc.; R7 = alkyl, -C(O)alkyl, alkoxyalkyl, etc.; p = 0-2; n = 0-2] or their salts were prepared For example, reaction of 3-([(1S)-2-[(difluoromethyl)oxy]-1-methylethyl]oxy)-5-hydroxy-N-(1-methyl-1H-pyrazol-3-yl)benzamide, e.g., prepared from Me 3,5-dihydroxybenzoate in 9 steps, with 1-(3-chloro-4-fluorobenzoyl)azetidine afforded compound II. Compds. of the invention generally activated glucokinase with an EC50 of less than about 500 nM, e.g., compound II exhibited the EC50 value of 40 nM.

ACCESSION NUMBER: 2007:61636 CAPLUS
DOCUMENT NUMBER: 146:142648
TITLE: Preparation of heteroaryl benzamide derivatives as glucokinase activators for the treatment of diabetes
INVENTOR(S): McKerrecher, Darren; Pike, Kurt Gordon; Waring, Michael James
PATENT ASSIGNEE(S): Astrazeneca AB, Swed.; Astrazeneca UK Limited
SOURCE: PCT Int. Appl., 62pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|----------|------------------|------------|
| WO 2007007042 | A1 | 20070118 | WO 2006-GB2472 | 20060603 |
| W: | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KN, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW | | | |
| RW: | AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | |
| JP 2009500444 | T | 20090108 | JP 2008-520937 | 20060703 |
| IN 2007DN10164 | A | 20080620 | IN 2007-DN10164 | 20071228 |
| CN 101218230 | A | 20080709 | CN 2006-80024889 | 20080108 |
| US 20080234273 | A1 | 20080925 | US 2008-995079 | 20080606 |
| PRIORITY APPLN. INFO.: | | | GB 2005-14174 | A 20050709 |
| | | | GB 2005-16298 | A 20050809 |
| | | | WO 2006-GB2472 | W 20060603 |

OTHER SOURCE(S): CASREACT 146:142648; MARPAT 146:142648

IT 919492-72-1P

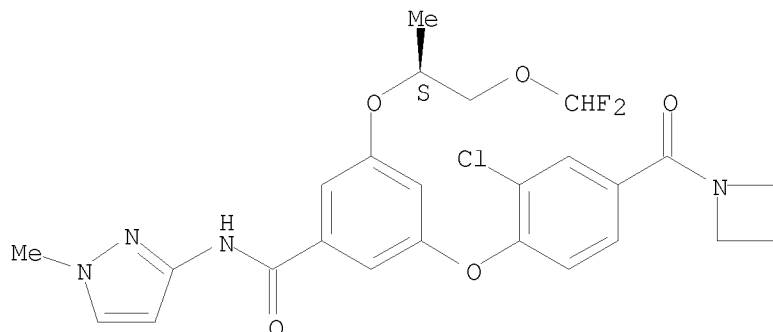
RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)

(preparation of heteroaryl benzamide derivs. as glucokinase activators for treatment of diabetes)

RN 919492-72-1 CAPLUS

CN Benzamide, 3-[4-(1-azetidinylicarbonyl)-2-chlorophenoxy]-5-[(1S)-2-(difluoromethoxy)-1-methylethoxy]-N-(1-methyl-1H-pyrazol-3-yl)- (CA INDEX NAME)

Absolute stereochemistry.



IT 919492-73-2P

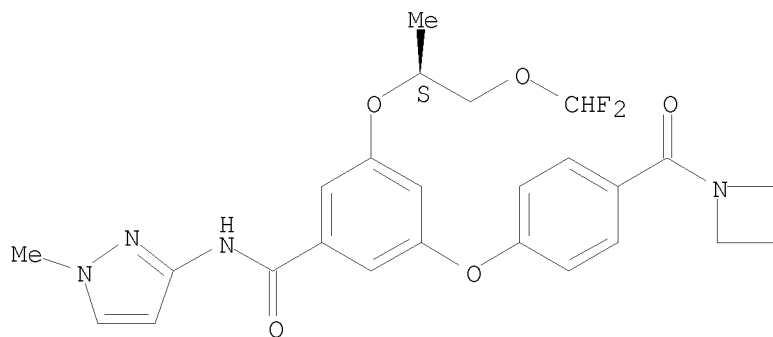
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of heteroaryl benzamide derivs. as glucokinase activators for treatment of diabetes)

RN 919492-73-2 CAPLUS

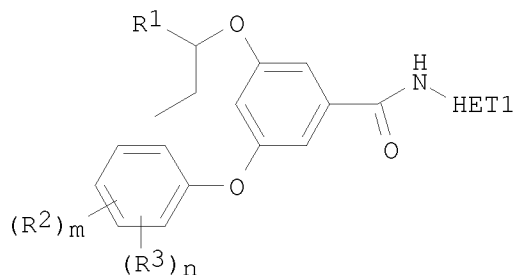
CN Benzamide, 3-[4-(1-azetidinylicarbonyl)phenoxy]-5-[(1S)-2-(difluoromethoxy)-1-methylethoxy]-N-(1-methyl-1H-pyrazol-3-yl)- (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 3 OF 4 CAPLUS COPYRIGHT 2009 ACS on STN
GI



I

AB Title compds. [I; R1 = MeOCH₂; R2 = CONR₄R₅, SO₂NR₄R₅, SOpR₄, HET₂; HET₁ = 5-6 membered (substituted) C-linked heteroaryl; HET₂ = 4-6 membered, C- or N-linked (substituted) heterocyclyl; R₃ = halo, FCH₂, F₂CH, CF₃, Me, MeO, cyano; R₄ = H, (substituted) alkyl, HET₂; R₅ = H, alkyl; R₄R₅N = HET₃; HET₃ = (substituted) N-linked, 4-6 membered, saturated or partially unsatd. heterocyclyl; n, p = 0-2; m = 0, 1; provided that when m = 0, then n = 1, 2], were prepared Thus, 3-[4-(azetidin-1-ylcarbonyl)-2-fluorophenoxy]-5-[[(1S)-1-(methoxymethyl)propyl]oxy]-N-(1-methyl-1H-pyrazol-3-yl)benzamide (preparation outlined) activated glucokinase with EC₅₀ = 0.04 μ M.

ACCESSION NUMBER: 2006:366940 CAPLUS

DOCUMENT NUMBER: 144:412497

TITLE: Preparation of N-pyrazolyl phenoxybenzamides as glucokinase activators for the treatment of type 2 diabetes.

INVENTOR(S): Johnstone, Craig; McKerrecher, Darren; Pike, Kurt Gordon; Waring, Michael James

PATENT ASSIGNEE(S): Astrazeneca AB, Swed.; Astrazeneca UK Limited

SOURCE: PCT Int. Appl., 87 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
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| WO 2006040528 | A1 | 20060420 | WO 2005-GB3888 | 20051011 |
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| CN 101039915 | A | 20070919 | CN 2005-80035074 | 20051011 |
| EP 1856056 | A1 | 20071121 | EP 2005-791490 | 20051011 |
| R: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LI, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR | | | | |
| JP 2008516936 | T | 20080522 | JP 2007-536248 | 20051011 |
| IN 2007DN02527 | A | 20070803 | IN 2007-DN2527 | 20070404 |
| US 20080280874 | A1 | 20081113 | US 2008-665163 | 20080605 |
| PRIORITY APPLN. INFO.: | | | GB 2004-23043 | A 20041016 |
| | | | WO 2005-GB3888 | W 20051011 |

OTHER SOURCE(S): MARPAT 144:412497

IT 883749-41-5P 883749-42-6P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

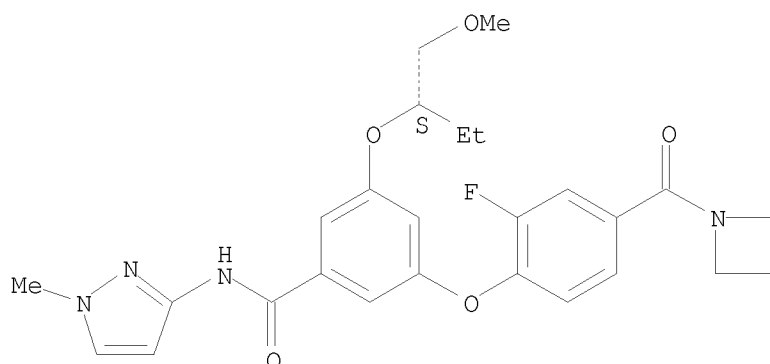
(claimed compound; preparation of N-pyrazolyl phenoxybenzamides as glucokinase

activators for the treatment of type 2 diabetes)

RN 883749-41-5 CAPLUS

CN Benzamide, 3-[4-(1-azetidinylicarbonyl)-2-fluorophenoxy]-5-[(1S)-1-(methoxymethyl)propoxy]-N-(1-methyl-1H-pyrazol-3-yl)- (CA INDEX NAME)

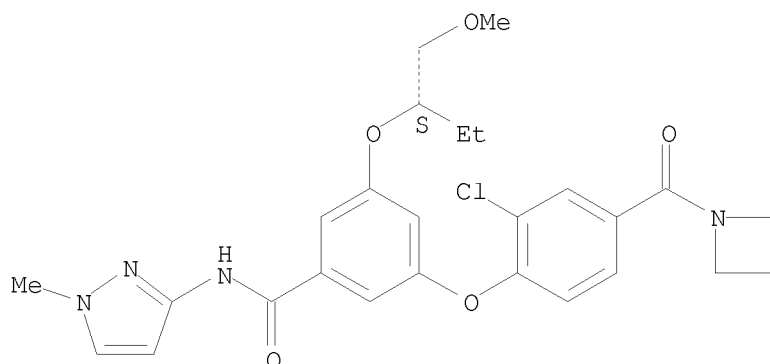
Absolute stereochemistry.



RN 883749-42-6 CAPLUS

CN Benzamide, 3-[4-(1-azetidinylicarbonyl)-2-chlorophenoxy]-5-[(1S)-1-(methoxymethyl)propoxy]-N-(1-methyl-1H-pyrazol-3-yl)- (CA INDEX NAME)

Absolute stereochemistry.



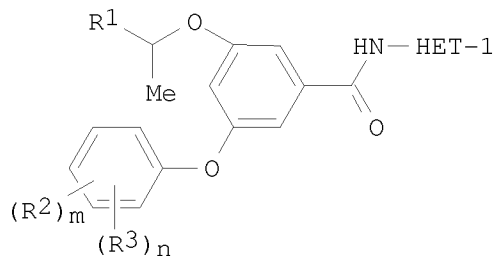
REFERENCE COUNT:

6

THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 4 OF 4 CAPLUS COPYRIGHT 2009 ACS on STN

GI



I

AB Title compds. I [R1 = methoxymethyl; R2 = carboxamido, sulfonamido, etc.; HET-1 = 5-6 membered C-linked heteroaryl; R3 = halo, fluoromethyl, difluoromethyl, etc.; m = 0-1; n = 0-2] are prepared For instance, 3-[4-[[[(2-methoxyethyl)amino]carbonyl]phenoxy]-5-[(1S)-2-methoxy-1-methylethyl)oxy]-N-(thiazol-2-yl)benzamide is prepared by the coupling of 4-[[3-[[[(1S)-2-methoxy-1-methylethyl)oxy]-5-[(thiazol-2-ylamino)carbonyl]phenyl]oxy]benzoic acid (preparation given) and 2-methoxyethylamine (DMF, DIPEA, HATu). Compds. of the invention generally have an activating activity for glucokinase with an EC50 of < 500 nM. I are useful in the treatment of type 2 diabetes.

ACCESSION NUMBER: 2005:962230 CAPLUS
DOCUMENT NUMBER: 143:266914
TITLE: Preparation of N-heteroaryl aryloxy-substituted benzamides as glucokinase activating agents
INVENTOR(S): Johnstone, Craig; McKerrecher, Darren; Pike, Kurt Gordon
PATENT ASSIGNEE(S): Astrazeneca AB, Swed.; Astrazeneca UK Limited
SOURCE: PCT Int. Appl., 138 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:

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| WO 2005080359 | A1 | 20050901 | WO 2005-GB545 | 20050215 |
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| RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| AU 2005214132 | A1 | 20050901 | AU 2005-214132 | 20050215 |
| CA 2554310 | A1 | 20050901 | CA 2005-2554310 | 20050215 |
| EP 1718624 | A1 | 20061108 | EP 2005-708360 | 20050215 |
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| CN 1922159 | A | 20070228 | CN 2005-80005262 | 20050215 |
| BR 2005007746 | A | 20070710 | BR 2005-7746 | 20050215 |
| JP 2007523142 | T | 20070816 | JP 2006-553652 | 20050215 |
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| US 20080280872 | A1 | 20081113 | US 2006-588334 | 20060803 |
| MX 2006PA09511 | A | 20061107 | MX 2006-PA9511 | 20060818 |
| KR 2007007103 | A | 20070112 | KR 2006-719123 | 20060918 |
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| | | | GB 2004-23039 | A 20041016 |
| | | | WO 2005-GB545 | W 20050215 |

OTHER SOURCE(S): CASREACT 143:266914; MARPAT 143:266914

IT 863504-11-4P 863504-45-4P 863504-46-5P
863504-47-6P

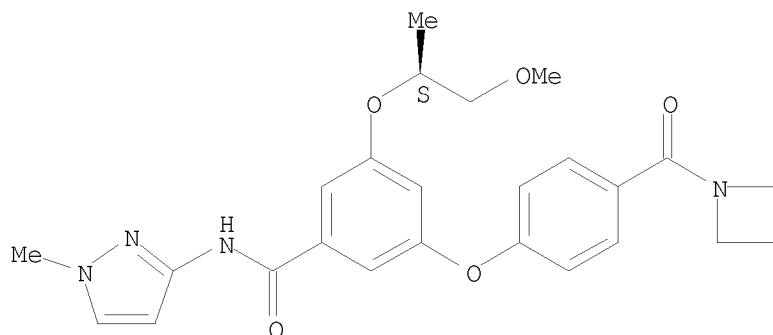
RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU
(Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
(Uses)

(preparation of N-heteroaryl aryloxy-substituted benzamides as glucokinase
activating agents)

RN 863504-11-4 CAPLUS

CN Benzamide, 3-[4-(1-azetidinylicarbonyl)phenoxy]-5-[(1S)-2-methoxy-1-
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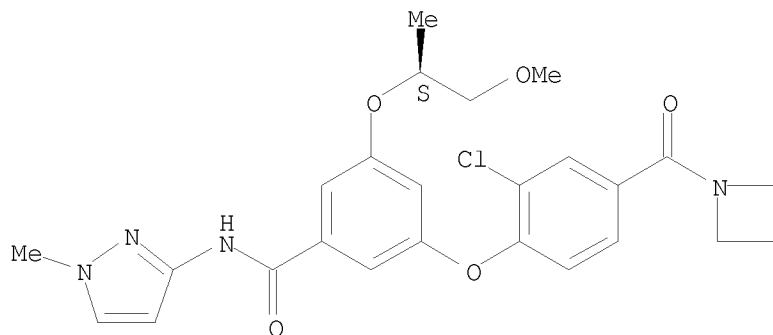
Absolute stereochemistry.



RN 863504-45-4 CAPLUS

CN Benzamide, 3-[4-(1-azetidinylicarbonyl)-2-chlorophenoxy]-5-[(1S)-2-methoxy-
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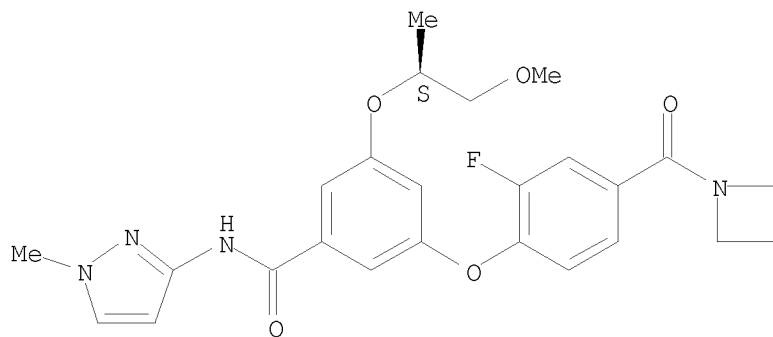
Absolute stereochemistry.



RN 863504-46-5 CAPLUS

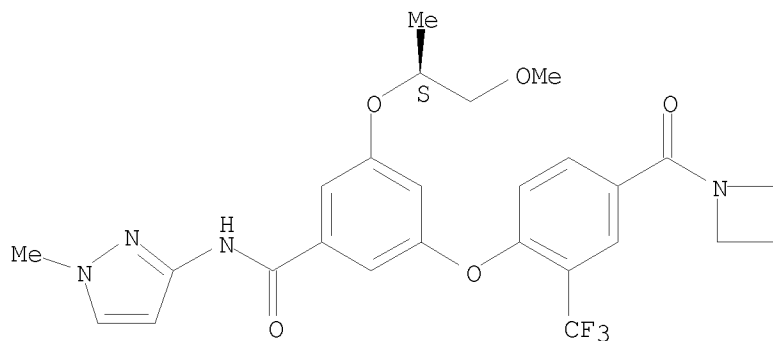
CN Benzamide, 3-[4-(1-azetidinylicarbonyl)-2-fluorophenoxy]-5-[(1S)-2-methoxy-
1-methylethoxy]-N-(1-methyl-1H-pyrazol-3-yl)- (CA INDEX NAME)

Absolute stereochemistry.



RN 863504-47-6 CAPLUS
 CN Benzamide, 3-[4-(1-azetidiny carbonyl)-2-(trifluoromethyl)phenoxy]-5-[(1S)-2-methoxy-1-methylethoxy]-N-(1-methyl-1H-pyrazol-3-yl)- (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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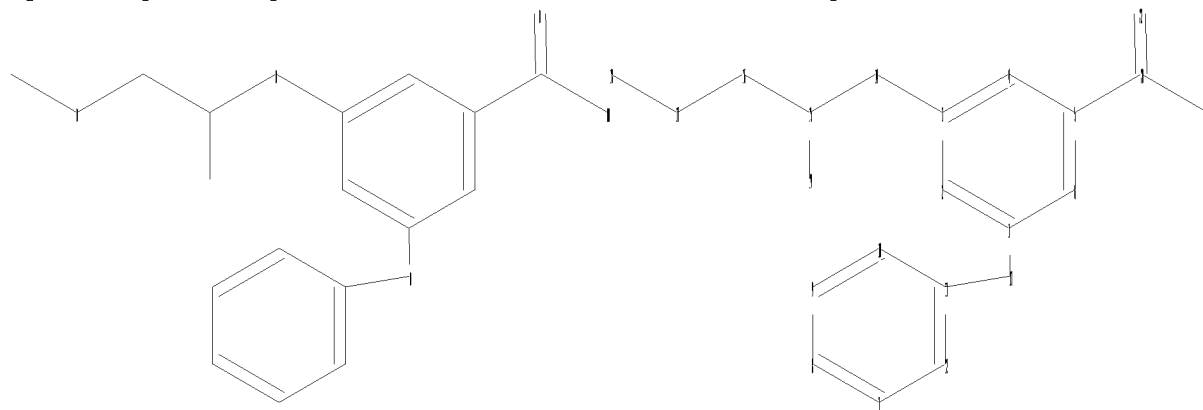
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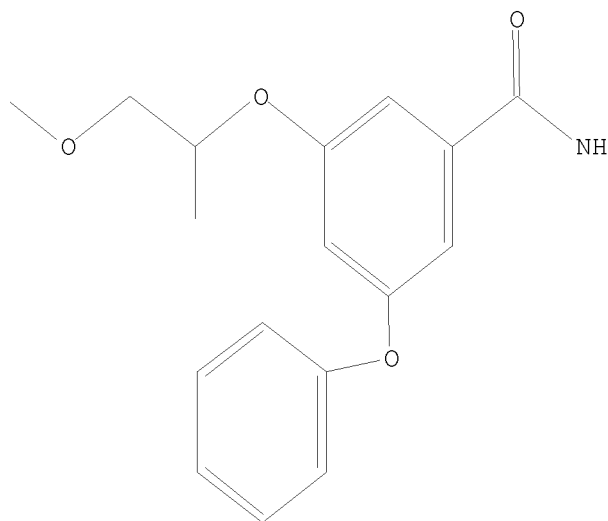
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FILE LAST UPDATED: 20 Jan 2009  (20090120/ED)
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4502964 AY<2003

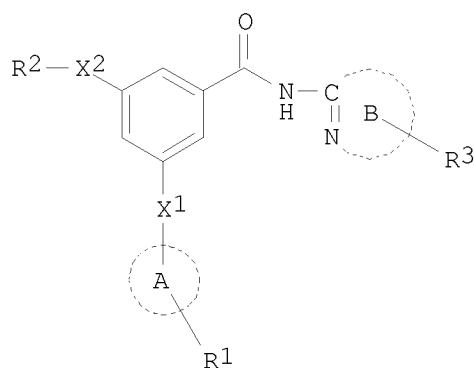
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L4 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2009 ACS on STN

GI



AB The title compds. (I) [wherein X1 =O, S, NH; X2 = O, S, CH2; R1 = 1 or 2 groups selected alkylsulfonyl, alkanoyl, lower alkyl, hydroxyalkyl, HO, mono or dialkylcarbamoyl, mono- or dialkylsulfamoyl, alkylthio, alkoxy, alkoxy-carbonylamino, alkoxy-carbonyl, halo, alkanoylaminoalkyl, alkoxy-carbonylaminoalkyl, alkylsulfonylaminoalkyl, cyano, and CF3 on the ring A; R2 = (un)substituted C3-7 cyclic alkyl optionally having one of the carbon atoms (excluding the carbon linked to X2) on the ring replaced by O, NH, N-alkanoyl, or NHC=O; R3 = 1 or 2 groups selected from lower alkyl, alkoxy, mono- or dialkylamino, halo, CF3, hydroxyalkyl, alkoxyalkyl, aminoalkyl, alkanoyl, CO2H, alkoxy-carbonyl, and cyano on the ring B; the ring A = 6- to 10-membered aryl or 5- to 7-membered heteroaryl; the ring B = mono- or bicyclic heteroaryl wherein the carbon atom bonded to the amide N atom forms C:N together with the ring N atom] or pharmaceutically acceptable salts thereof. These compds. and salts thereof function to activate glucokinase and are useful as preventive and/or therapeutic agents for obesity or diabetes or for the treatment, prevention, and/or onset of type II diabetes. Thus, S-oxidation of 5-[[[(1S)-2-(tert-butyl-dimethylsilyloxy)-1-methylethyl]oxy]-3-(4-methylthiophenoxy)benzoic acid Me ester by m-chloroperbenzoic acid in CHCl3 under ice-cooling, saponification with a mixture of 5 N NaOH and MeOH and acidification with 5% citric acid, amidation with 2-aminothiazole using 1-ethyl-3-(3-dimethylaminopropyl)carbodiimide hydrochloride and 1-hydroxybenzotriazole hydrate in CH2Cl2, and finally desilylation with 4 N HCl/dioxane at room temperature for 15 min gave 5-(2-hydroxy-1-methylethoxy)-3-[4-(methanesulfonyl)phenoxy]-N-(thiazol-2-

yl)benzamide (II). II activated recombinant human liver glucokinase with
EC50 of 0.08 μ M.

ACCESSION NUMBER: 2004:740302 CAPLUS

DOCUMENT NUMBER: 141:260754

TITLE: Preparation of heteroarylcarbamoylebenzene derivatives
as glucokinase activators

INVENTOR(S): Iino, Tomoharu; Hashimoto, Noriaki; Nakashima,
Hiroshi; Takahashi, Keiji; Nishimura, Teruyuki; Eiki,
Junichi

PATENT ASSIGNEE(S): Banyu Pharmaceutical Co., Ltd., Japan

SOURCE: PCT Int. Appl., 288 pp.
CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
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| WO 2004076420 | A1 | 20040910 | WO 2004-JP2284 | 20040226 <-- |
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| RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| AU 2004215514 | A1 | 20040910 | AU 2004-215514 | 20040226 <-- |
| CA 2516407 | A1 | 20040910 | CA 2004-2516407 | 20040226 <-- |
| EP 1600442 | A1 | 20051130 | EP 2004-714930 | 20040226 |
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| BR 2004007810 | A | 20060301 | BR 2004-7810 | 20040226 |
| CN 1777589 | A | 20060524 | CN 2004-80010759 | 20040226 |
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| ZA 2005005982 | A | 20060726 | ZA 2005-5982 | 20050726 |
| IN 2005DN03604 | A | 20070601 | IN 2005-DN3604 | 20050816 |
| US 20060167053 | A1 | 20060727 | US 2005-546962 | 20050825 |
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| | | | JP 2003-400882 | A 20031128 |
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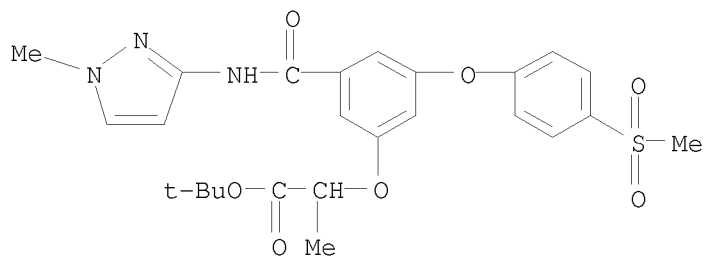
OTHER SOURCE(S): MARPAT 141:260754

IT 752240-23-6P

RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent); USES (Uses)
(preparation of heteroarylcarbamoylebenzene derivs. as glucokinase activators for treatment of diabetes)

RN 752240-23-6 CAPLUS

CN Propanoic acid, 2-[3-[[[(1-methyl-1H-pyrazol-3-yl)amino]carbonyl]-5-[4-(methylsulfonyl)phenoxy]phenoxy]-, 1,1-dimethylethyl ester (CA INDEX NAME)



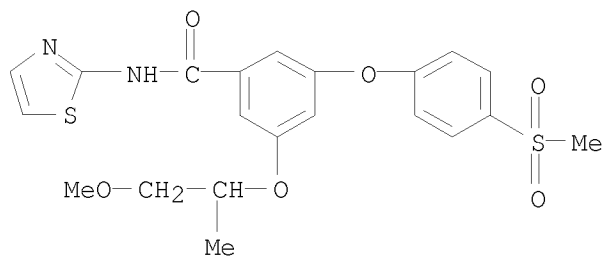
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 752239-74-0P 752239-98-8P 752240-73-6P
 752240-79-2P 752240-95-2P 752241-29-5P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU
 (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES
 (Uses)

(preparation of heteroarylcarbamoylbenzene derivs. as glucokinase activators
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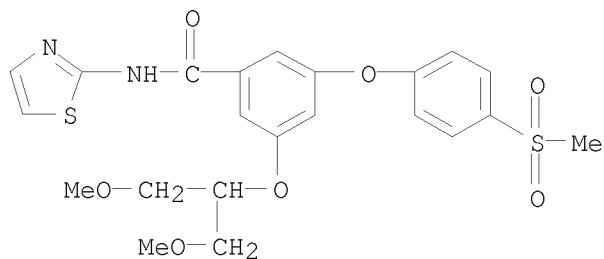
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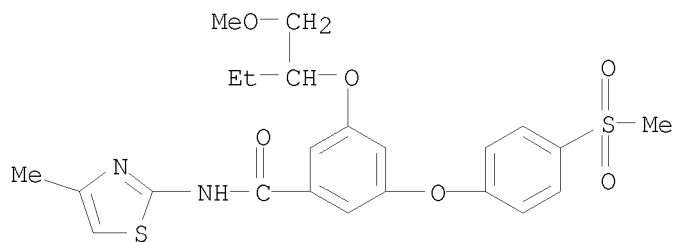
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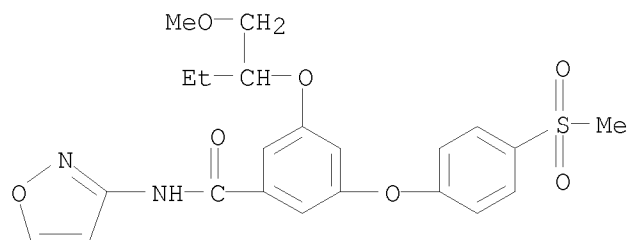
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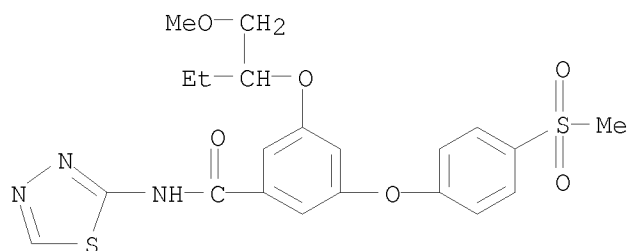
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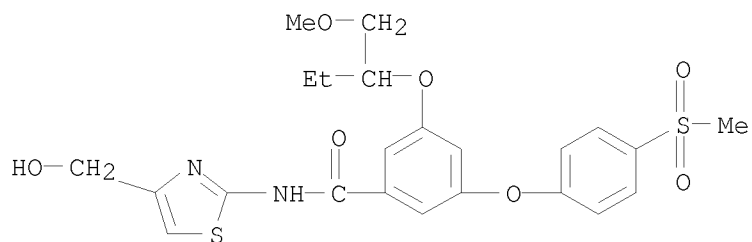
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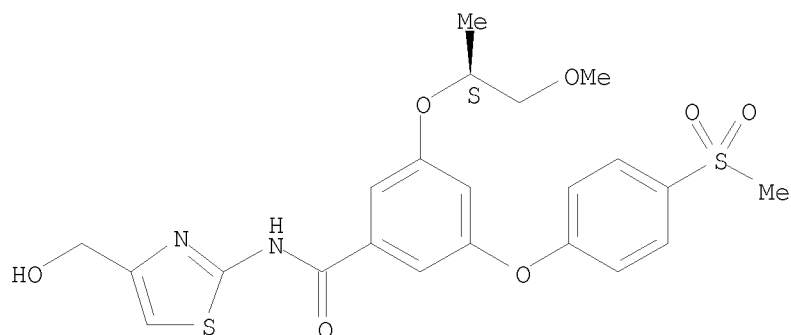
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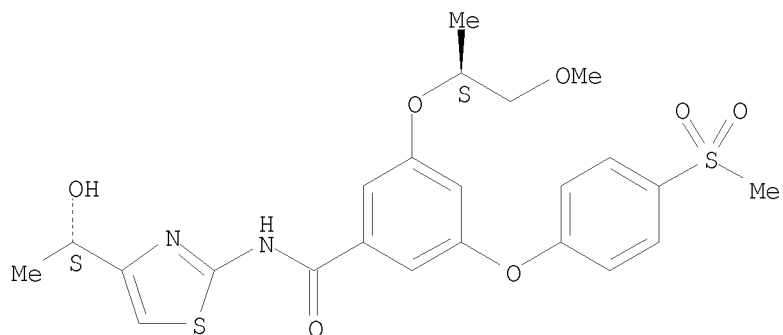
Absolute stereochemistry.



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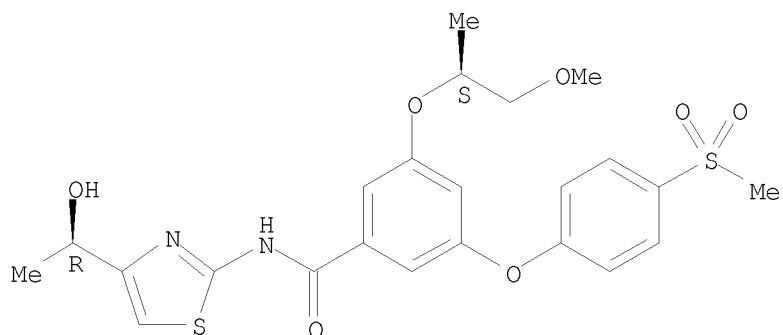
Absolute stereochemistry.



RN 752239-39-7 CAPLUS

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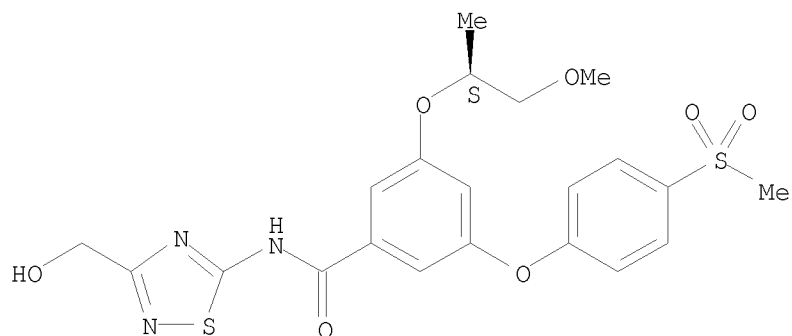
Absolute stereochemistry.



RN 752239-74-0 CAPLUS

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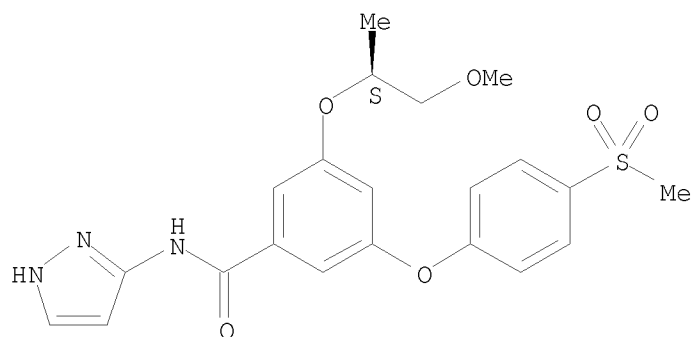
Absolute stereochemistry.



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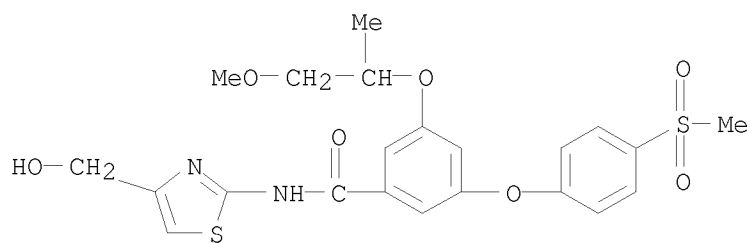
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Absolute stereochemistry.



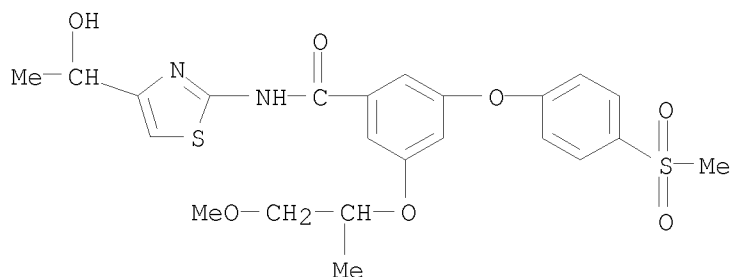
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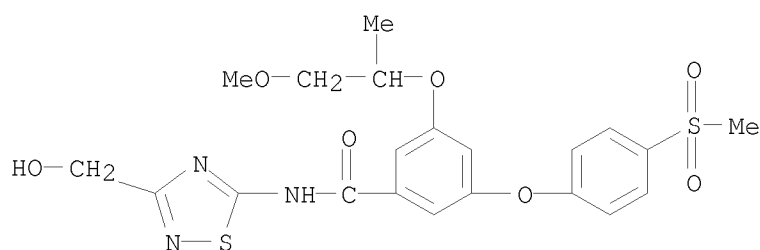
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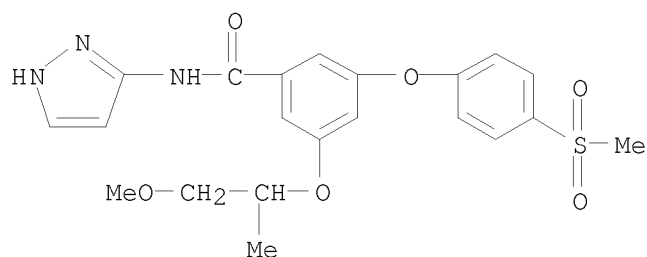
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RN 752241-29-5 CAPLUS

CN Benzamide, 3-(2-methoxy-1-methylethoxy)-5-[4-(methylsulfonyl)phenoxy]-N-1H-pyrazol-3-yl- (CA INDEX NAME)



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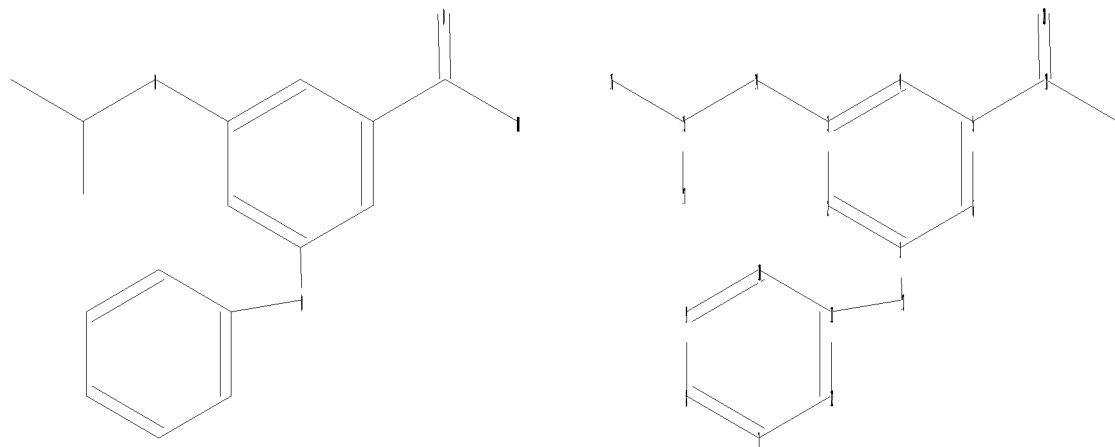
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<http://www.cas.org/support/stngen/stndoc/properties.html>

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Uploading C:\Program Files\STNEXP\Queries\10588334 generic2.str



chain nodes :
13 14 15 16 17 18 19 20
ring nodes :
1 2 3 4 5 6 7 8 9 10 11 12
chain bonds :
1-13 3-14 5-18 11-13 14-15 15-16 15-17 18-19 18-20
ring bonds :
1-2 1-6 2-3 3-4 4-5 5-6 7-8 7-12 8-9 9-10 10-11 11-12
exact/norm bonds :
1-13 3-14 11-13 14-15 18-19 18-20
exact bonds :
5-18 15-16 15-17
normalized bonds :
1-2 1-6 2-3 3-4 4-5 5-6 7-8 7-12 8-9 9-10 10-11 11-12

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom
11:Atom 12:Atom 13:CLASS 14:CLASS 15:CLASS 16:CLASS 17:CLASS 18:CLASS
19:CLASS 20:CLASS

L5 STRUCTURE UPLOADED

=> s l5 sss ful
FULL SEARCH INITIATED 13:17:53 FILE 'REGISTRY'
FULL SCREEN SEARCH COMPLETED - 4040 TO ITERATE

100.0% PROCESSED 4040 ITERATIONS 669 ANSWERS
SEARCH TIME: 00.00.01

L6 669 SEA SSS FUL L5

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| COST IN U.S. DOLLARS | SINCE FILE | TOTAL |
| | ENTRY | SESSION |
| FULL ESTIMATED COST | 185.88 | 385.84 |
| DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) | SINCE FILE | TOTAL |
| | ENTRY | SESSION |
| CA SUBSCRIBER PRICE | 0.00 | -0.82 |

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FILE COVERS 1907 - 21 Jan 2009 VOL 150 ISS 4
FILE LAST UPDATED: 20 Jan 2009 (20090120/ED)

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L7 17 L6

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25138568 PY<2005
4502964 AY<2003
3971717 PRY<2003
L8 1 L7 AND (PY<2005 OR AY<2003 OR PRY<2003)

=> d 18

L8 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2009 ACS on STN
AN 2004:740302 CAPLUS
DN 141:260754
TI Preparation of heteroarylcarbamoylbenzene derivatives as glucokinase
activators
IN Iino, Tomoharu; Hashimoto, Noriaki; Nakashima, Hiroshi; Takahashi, Keiji;
Nishimura, Teruyuki; Eiki, Junichi
PA Banyu Pharmaceutical Co., Ltd., Japan
SO PCT Int. Appl., 288 pp.
CODEN: PIXXD2
DT Patent
LA Japanese
FAN.CNT 1

| | PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------|----------|------------------|--------------|
| PI | WO 2004076420 | A1 | 20040910 | WO 2004-JP2284 | 20040226 <-- |
| | W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI | | | | |
| | RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| | AU 2004215514 | A1 | 20040910 | AU 2004-215514 | 20040226 <-- |
| | CA 2516407 | A1 | 20040910 | CA 2004-2516407 | 20040226 <-- |
| | EP 1600442 | A1 | 20051130 | EP 2004-714930 | 20040226 |
| | R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK | | | | |
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| | ZA 2005005982 | A | 20060726 | ZA 2005-5982 | 20050726 |
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| | US 7432287 | B2 | 20081007 | | |
| | NO 2005004425 | A | 20050923 | NO 2005-4425 | 20050923 |
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| PRAI | JP 2003-49466 | A | 20030226 | | |
| | JP 2003-400882 | A | 20031128 | | |
| | JP 2004-31298 | A | 20040206 | | |
| | WO 2004-JP2284 | A | 20040226 | | |
| | US 2005-546962 | A3 | 20050825 | | |

OS MARPAT 141:260754

RE.CNT 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD
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COST IN U.S. DOLLARS

| SINCE FILE | TOTAL |
|------------|---------|
| ENTRY | SESSION |
| 8.47 | 394.31 |

FULL ESTIMATED COST

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

| SINCE FILE | TOTAL |
|------------|---------|
| ENTRY | SESSION |
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